

Amendments to the Specification:

Please replace the paragraph beginning on page 64, line 33, with the following rewritten paragraph:

--Pharmacologically, α_2 -adrenergic receptors are defined as highly sensitive to the antagonists yohimbine ($K_i = 0.5\text{-}25 \mu\text{M}$), atipamezole ($K_i = 0.5\text{-}2.5 \mu\text{M}$), and idazoxan ($K_i = 21\text{-}35 \text{ nM}$) and with low sensitivity to the α_1 receptor antagonist prazosin. Agonists selective for the α_2 -adrenergic receptor class relative to the α_1 -adrenergic receptor class are UK14304, BHT920 and BHT933. Oxymetazoline binds with high affinity and selectivity to the α_{2A} -receptor subtype ($K_D = 3 \mu\text{M}$), but in addition binds with high affinity to α_1 -adrenergic receptors and 5HT1 receptors. An additional complicating factor is that α_2 -adrenergic receptor ligands which are imidazolines (clonidine, idazoxan) and others (oxymetazoline and UK14304) also bind with high affinity (nanomolar) to non-adrenoceptor imidazoline binding sites. Furthermore, species variation in the pharmacology of the α_{2A} -adrenoceptor exists. To date, subtype-selective α_2 -adrenergic receptor ligands show only minimal selectivity or are nonselective with respect to other specific receptors, such that the therapeutic properties of subtype selective drugs are still under development.--

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